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IN STUDIES OF ITS CONTEMPORARY FORMS

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- USSR -

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THE TYPHUS FEVER PROBLEM AS REFLECTED  
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-USSR-

[Following is the translation of an article entitled "Problema sypnogo tifa v otrazhenii izucheniya yego sovremennykh form" (English version above) by Prof P. F. Zdrodovskiy in Vestnik Akademii Meditsinskikh Nauk SSSR (Herald of the Academy of Medical Sciences USSR), Vol 15, No 7, Moscow, 1960, pages 3-14.]

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Among the peculiarities of residual forms of contemporary typhus there are included: the sporadic character of the diseases observed, as a rule without the detection of the disease sources and in the absence in the foci of carriers of the infection -- body lice; the primary concentration (in certain localities up to 80%) of the disease in the older age group (41-60 years of age and over) in contrast to past times when, for the most part (45-62%) typhus affected people in the young and mature years (20-40 years of age); there is an unusual number of recurrent attacks of typhus (40% and higher); there is the benign character of the disease with the extremely rare fatal outcome; there is the absence or obliteration of the winter-spring seasonal prevalence so characteristic of typhus in the past. To the above should be added the relative difficulty of isolating from the blood of the diseased the pathogenic agent which retains all the visual characteristics of Provachek's rickettsia, i.e., of the authentic stimulus of typhus. It is impossible, likewise, to ignore the limited sensitivity of the Weil-Felix reaction in the contemporary forms of typhus, which forms are at the same time serologically well differentiated when Provachek's rickettsia antigen is used. These characteristics of the major portion of contemporary forms of typhus are

responsible for

^disorienting, in no small degree, epidemiologists, who from established tradition continue the analysis of these forms from epidemiological points of view, which have been developed for typhus.

As is known, the code of epidemiology, having become established in the minds of the majority of epidemiologists, runs as follows: the sole source of infection in typhus is the human being ill with the typhus; the sole carrier of this infection is the louse (primarily, the body louse), which becomes diseased in sucking the blood of the diseased person. Aside from the epidemiological chain "louse-human-louse", infection with exanthematous fever (or typhus) cannot occur in natural circumstances. To the facts indicated, established half a century ago by C. Nicolle unquestionably for epidemic typhus, there is being added the absence in typhus fever both of the "obliterated" and "asymptomatic" forms of the infection by the majority of our epidemiologists, led by L. V. Gromashevskiy.

In this direction L. V. Gromashevskiy writes in his work on local epidemiology (1947): "In diseases, which exhibit a wide polymorphic range in the clinical appraisal, proceeding in the direction of increasingly mildly progressing ("obliterated") forms, it may be expected that such conditions will be met when every reaction to the infection from the part of the organism is absent; such a condition is designated as a healthy carrier state (infection carrier state). And, on the contrary, in infections the course of which always progresses in a clinically manifest form and do not give the "obliterated progression", the possibilities of an infection carrier state are excluded" (page 424). Further, proceeding from his numerous observations that the length of the fever in typhus is never less than nine days -- this is the minimum period, which accounts for only 2.1 % of the disease -- L. V. Gromashevskiy concludes: "This most important fact for the clinic (diagnostics) and epidemiology denies with a certitude the possibility of the existence of a healthy carrier state in typhus, for a disease in which the mildest observed forms proceed as a nine-day fever cannot proceed as an asymptomatic healthy carrier state" (page 421).

The indicated thesis of L. V. Gromashevskiy entered the official code of epidemiology for typhus in its entirety, having been confirmed in 1939 by the All-Union Conference of Microbiologists, Epidemiologists,

and Infectious Disease Specialists. As a result of the report by L. V. Gromashevskiy, "the recently prevailing teaching of virus-carrying in typhus, in the literature of our country" was admitted as groundless, and the estimate of the epidemiological role of this factor was found to be based on "no concrete data whatsoever" (quoted from the resolution of the conference). Since then, the question of obliterated forms and virus-carrying in typhus has been relegated to the category of harmful epidemiological mistakes, deserving of the most severe judgment.

Occasioned by the conflict which had occurred between the prevailing ideas of the epidemiology of typhus and the real state of affairs, and also from considering the data of world-wide experimentation in the field of rickettsial disease and, in part, also in typhus, we decided to subject some of the collected dogmas in the teaching of this infection to revision. The need for such a revision was supported by the fact that the existing ideas on the epidemiology of typhus, as has already been indicated above, came to our attention as a result of the analysis of epidemic forms of this disease. Also, the study of typhus had previously been carried out with primitive methods of research, while contemporary science makes use of methods which are further perfected, specifically, the method of the highly effective specific flow and the retrospective serodiagnosis aided by the reactions of complement fixation (RSK) [reaktsiya svyazyvaniya komplementa - complement fixation reaction] with the use of antigen from cultures of Provanchek's rickettsia.

At the first stage we examined the thesis indicated above of the absence in typhus of "obliterated" and "asymptomatic" forms of the infection. This thesis, in spite of the existence of sufficiently well-founded objections from some Soviet investigators (K. R. Tokarevich and colleagues), continues to prevail among our epidemiologists, in large measure determining their tactics relating to typhus, and so it demands therefore an immediate resolution in one direction or the other.

We placed before ourselves the problem of instituting a mass serological investigation of population groups in various regions, which groups have been infected in the past, to one degree or another, with typhus, and paying special attention to the age distribution of the results of serological researches.

The study of typhus was carried out by immunological methods of retrospective diagnostics, with both wide and narrow application in virus infection (poliomyelitis, encephalitis, and others) and in a number of bacterial infections (diphtheria, scarlet fever, and others).

We organized, in 1957, in cities of the central region of the RSFSR corresponding investigations with mass use of contemporary methods of serodiagnostics of typhus infection with the aid of RSK. Designed under one plan, with one method of research, and with the use of one standard antigen for RSK, these observations during 1957 - 1958, and partly in 1959 were carried out on a mass scale by the efforts of the specially prepared scientific workers of local institutes of microbiology and epidemiology, sanitation - epidemiological stations and other laboratories under the general leadership of the author and under the methodological leadership of E. M. Golinevich. During 1957-1958 a collective composed of 30 scientific workers completed, essentially, the work assigned it on a series of questions, having developed a personal research initiative [See Note]. As a result there was collected unique material on the study of the epidemiology of typhus, the basic results of which are briefly summarized in this article.

(Note The authors of the research material summarized in this article are V. A. Yablonskaya, T. S. Kovreva, A. V. Yermenko (Moscow); N. I. Stratonovich, A. I. Yakushkina, L. A. Pravdina, G. N. Rozanova, A. I. Kogan, T. A. Manevich, Ye. S. Parfenova, Ye. S. Temryuk (Smolensk); G. R. Gazizova, A. E. Reznik, N. R. Bayteryskova, M. P. Lashmanova, F. B. Kolpachikhin, P. A. Sergeeva, V. M. Klyuchnikova (Kazan'); G. I. Grenaus, A. S. Antsunova, V. T. Degtyareva, Ye. A. Miloradovskaya, I. S. Soldatova, M. V. Lopatina (Gor'kiy); L. N. Astakhova, M. V. Matveyeva, G. P. Setnovskaya (Sverdlovsk); M. A. Kulemina, K. P. Kuznetsova (Kuybyshev). The material of the authors indicated is cited from their manuscripts which are subject to publication.)

A comparison of the objective indices of serological research with the data of anamnesis confirmed that in mass observations the anamnestic data display sufficient authenticity. Thus, among 5124 cases of serological investigation with negative typhus according to the RSK anamnesis, the RSK with the typhus antigen in 10 % of the diseased was found, on the average, to be positive, while among 362 persons with

positive typhus the RSK anamnesis was recorded to be 41 % on the average, i. e., four times more frequent [See Note]. We will note in passing that K. N. Tokarevich likewise confirms the good correlation of immunological data from observations with the data of typhus anamnesis.

(Note) The following data convincingly illustrate, at the same time, the RSK specificity.)

Before presenting the results obtained and their discussion, we will turn to the elucidation of some features concerning the methodology used in the serological investigations.

As has previously been indicated, serological investigations were conducted with the aid of RSK, the methodology of foreign countries paralleling the methodology accepted by our laboratory, and, specifically, parallel with the methodology which was used by Rumanian investigators (D. Combesco and colleagues).

As antigens in the RSK arrangement, the so-called completely soluble antigens suggested by Ye. M. Golinevich were used, which are prepared from hemolyzed egg cultures of Provachek's rickettsia (the same type of antigens were used by the Rumanian authors). The RSK was applied with four units of the antigen indicated and with two units of the serum complement -- by "hot" (36 to 37 °) or by "cold" (40 °) methods without noticeable difference in the results. In mass observations with the goal of retrospective diagnostics of typhus infection, the serums under test were usually employed in the following dilutions: 1:5, 1:10, 1:20, 1:40, etc. In this area, part of the authors considered as positive (+ + + +) at the 1:5 dilution, the other part of the authors -- at the 1:10 dilution.

Titers of the positive RSK and their correlations in mass investigations of a healthy population with negative anamnesis for typhus are illustrated by table 1, which sums up the results of serological investigations in four cities.

As seen in table 1, positive RSK in a serum dilution of 1:5, on the average, gave us a registration of 18 %. How can the significance of this trial using the 1:5 dilution be evaluated? The Rumanian authors headed by Combesco (1957, 1958), having studied the significance of various RSK titers by a methodology paralleling ours, which is one of diagnostic titers for typhus, accepted the dilution 1:16 on the condition

that it be dynamically raised or lowered with repeated investigations. Parallel with this in their data, pos-

TABLE ONE

Titers of Positive RSK in Healthy People with  
Negative Anamnesis for Typhus

Number Investigated	Number of RSK Positive	Distribution of RSK Titers in Percent					
		1:5	1:10	1:20	1:40	1:80	1:160
3,257	330	11	18	46	26.4	7.3	2.3

itive RSK figures in the 1:4 dilution observed, on the one hand, at the beginning of typhus attacks and, from the other, in cases of late retrospective diagnostics of typhus infection in subjects with medical documentation of previous typhus experience. In this manner, in retrospective investigations a positive RSK at 1:5 dilution can specifically expose the early-experienced typhus infection.

Together with this, as special investigations of the Rumanian authors show, in separate cases it is impossible to exclude the unspecific RSK delay. Thus, in the deliberate investigation of 90 healthy children with negative anamnesis for typhus these authors once discovered a positive RSK at a 1:4 titer.

Concerning positive RSK with a 1:10 titer, we have no doubt of its specificity in retrospective diagnostics of typhus infection. Thus, from the data of N. I. Stratovich, R. I. Yakushkina, L. A. Pravdina, and others (Smolensk), among the 142 people investigated in the three to nineteen year range with deliberately negative typhus anamnesis, positive RSK was not observed in a single case with a serum dilution of 1:10.

We related to positive RSK, in the data presented for the mass retrospective investigation of population groups on typhus infection, the trials with 1:5 titer together with 1:10 trials, and higher dilutions. From this, as is seen from the figures below, we discovered no differences in the general regularity of the age distribution of positive RSK in the cases investigated, starting with a beginning titer of 1:5 (figure 2) or 1:10 (figure 1).

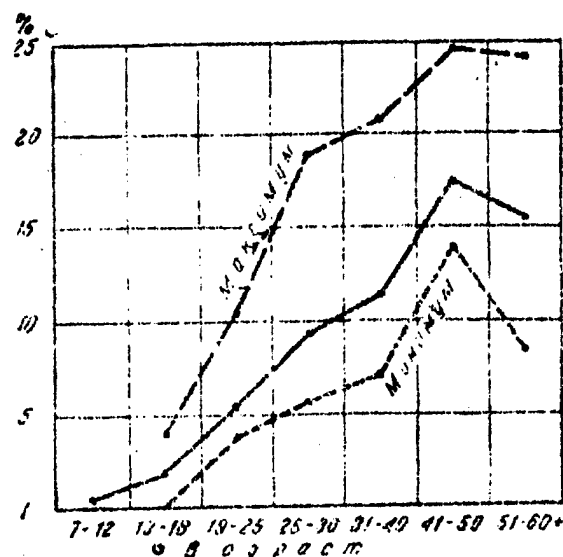


Figure 1. Summary curve of age distribution of positive RSK in population groups with negative anamnesis for typhus (in five cities of the RSFSR) [Upper dashed line reads "Maximum"; lower dashed line reads "Minimum." / Abscissa: Age]

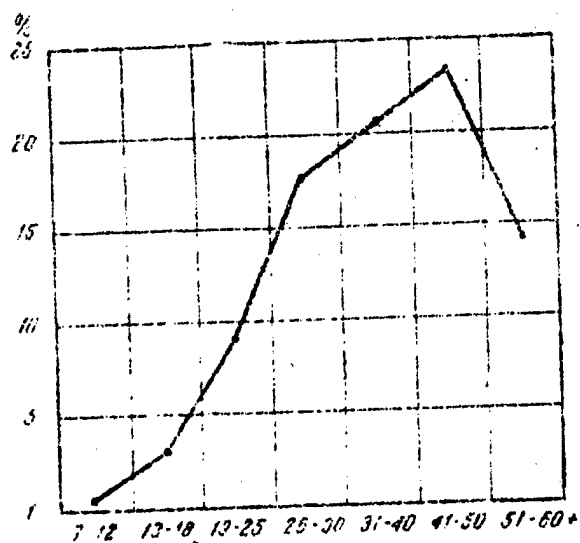


Fig. 2. Curve of the age distribution of positive RSK in population groups with negative anamnesis for typhus. Minimum RSK titer: 1:10 (City X) [Abscissa: Age]

Finally, for the character of general and particular RSK specificity with the antigen from Provachek's rickettsia it is necessary to indicate that V. A. Yablonskaya in our laboratory for about 50 cases of retrospective positive RSK to typhus infection carried out researches, using as controls the same serums for RSK, with the antigen from rickettsia of the tick-borne group, and obtained negative results.

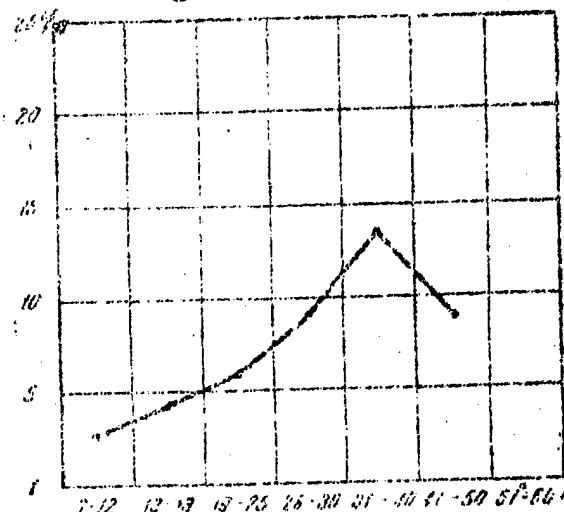


Figure 3. Age distribution curve of positive RSK in population groups with negative anamnesis to typhus. Minimum RSK titer: 1:5 to 1:10 (City S) / Abscissa: Age

The results of the retrospective serological RSK investigation of typhus infection of population groups with negative typhus anamnesis are presented in figure 1, where the positive RSK findings are summarized for the 5124 persons investigated in five cities of the central region of the RSFSR, with the distribution of positive RSK in various age groups.

In this manner, as is seen from figure 3, the summary curve of positive RSK progressively grows from zero or minimum importance in the child and in the younger ages to the mature and elderly age groups with the peak occurring at 41-50 years of age -- the type of curve is characteristic for summary data of population groups of five cities and regularly repeats for investigated population groups of different cities (figures 1, 2, and 4).

The trustworthiness of the data presented is supported by the presence of 80 % positive RSK results in titers which raise the minimum level of typhus infection in retrospective diagnosis. From the other side, serological data do not occasion any doubt, even in the statistical sense, since every age group, in the material summarized, is represented by highly significant figures (in the range of from 380 to 1156 persons to an age group).

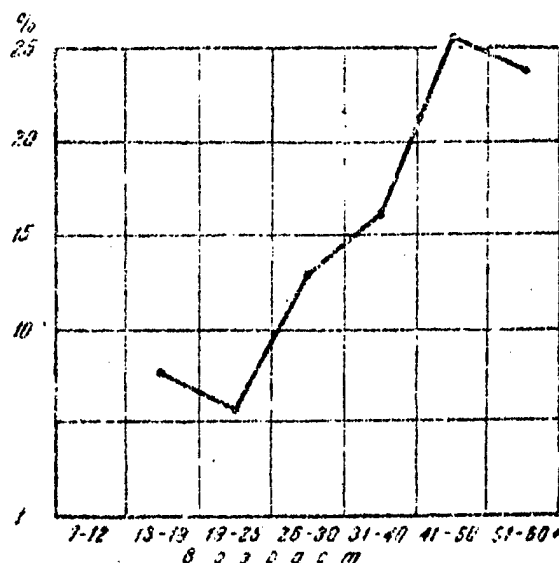


Figure 4. Age distribution curve of positive RSK in population groups with negative anamnesis for typhus (City K) [Abacissa: Age]

Analysis of the data discloses, first of all, the very low percentage of a positive RSK in the younger age groups (7-12 and 13-18 years of age). This circumstance indicates that in recent years the circulation of the typhus virus in the collectives investigated was absent or was insignificantly expressed in contrast to previous years (in particular, during the war period) when it was evidently important, as is reflected in the relatively high percentage of a positive RSK (11.7 to 17 % on the average) in the mature and elderly age groups (31-40 and 41-50 years of age and higher).

Simultaneously, in the 31-60 years of age and higher age group, with a categorical negation of previous typhus illness together with a negation, in general, of its occurrence in infection hospitals (see

figure 1), a positive RSK was, in places, registered at 21-25 %, i.e., was present in a very high percentage of the cases investigated (see figures 1 and 4). In connection with this it is necessary to note that the percentage displayed of a positive RSK is evidently understated, since sufficiently trustworthy evidence in the USSR and in foreign countries shows that in cases of clinically expressed typhus the positive RSK can, with time, disappear in a considerable percentage of those who were diseased.

As an illustration of what has been stated, we present the observations of M. A. Kulemnaya and K. P. Kuznetsova (Kuybyzhev), which, for 107 cases who experienced typhus of varying remoteness in time, there was determined the loss of a positive RSK, depending on the remoteness of the illness with the following correlations: for the course of one year RSK was retained by 38 of 41 of those who were diseased; for a period of 6-20 years by nine out of 21 persons; for a period of over 20 years by 16 out of 45 persons. Of 107 who had been ill in the periods indicated, a positive RSK loss occurred in 44 persons, or 41 %.

What conclusions should be drawn from the data presented of the mass serological investigation of population groups of older and elderly ages?

Considering the specificity of the RSK, it should be admitted that in the above-mentioned groups where the percentage of a positive RSK reached 21-25%, the investigated cases, in spite of their assertions, were ill in the 30-50 year age range with obvious forms of typhus, which seems absurd, or they were ill with typhus in some light "obliterated" form, having slipped away from recognition, which seems more reasonable and correlates with the observations in other countries (see below).

Direct observations on the length of the fever give a convincing illustration of the clinical polymorphism of contemporary forms of typhus in their general benign character distinguished universally.

As an illustration we present the investigations pertinent here, of M. A. Kulemina and K. P. Kuznetsova (Kuybyzhev), who studied the length of fever in 264 ill persons with typhus in contrast to the data of L. V. Gramashevskiy, relating to epidemic typhus (Table 2).

By analogy with the material indicated, according to the data of V. A. Yablonskaya, T. S. Kovreva and A. V. Yermanko (1956), among the 184 cases of

TABLE TWO

Length of Fever in Typhus (from data of native scientists)

Length of fever (in days)	Number of the diseased (in %)	
	From data of M. A. Kulemina and K. P. Kuznetsova (Kuybyshev)	From data of L. V. Groma- shevskiy
5-8	20	0
9	18.8	2.1
10-11	40	12.7
12-17	21.2	85.2

TABLE THREE

Length of Fever in Typhus (from the data of  
D. Combesco)

Length of fever (in days)	Number of the diseased (in percent)
4-6	6
7-9	48
10-11	28
12-14	18

sporadic typhus studied by them, the length of the fever oscillated within the range of from 6 to 15 days with an average length of 9.4 days. From the similar data of A. N. Astakhova, M. V. Matveyeva and G. P. Setunovskiy (1958), among 153 ill with typhus, the fever length was observed to be 7-9 days in 31.5 %, 10-12

days in 66 %, and 13-16 days in 6.9 % only (as against 85 % according to the data of L. V. Gramshevskiy).

In conclusion we present the data of D. Combesco and his colleagues (1957), concerning the length of fever of repetitive typhus in Rumania (table 3).

In other words, D. Combesco registered even a four-day fever length for typhus.

Thus, we see that in short-fever forms, the absence of which, in L. V. Gramshevskiy's investigations, is the main argument for the negation of "obliterated" and "asymptomatic" forms of typhus infection, at present are not rare -- rather, they are commonplace. It cannot be doubted that the clinical course of typhus has changed greatly in comparison with the past -- this disease has become considerably more benign, but its clinical expression more polymorphous, which, obviously, explained the difficulties observed in the clinical diagnosis of contemporary forms of typhus in the sporadic category and its frequent registration under an erroneous diagnosis.

In connection with this, the observations of M. A. Kulemina and K. P. Kuznetsova should be observed, who, in the investigation of 862 healthy subjects, retrospectively separated 142 persons with a positive RSK to typhus. It was found that among the latter, 63 persons or 44.3 % during the years 1957-1958 had influenza -- the diagnosis which sent such patients often and, evidently, not accidentally to the hospitals.

It should be said that in cases of hospitalization and lengthy observation not so rarely is it that contemporary typhus is clinically observed. Thus, among 832 cases of sporadic typhus studied, who had been hospitalized during the last few years in cities of the RSFSR, 75 persons, i.e., 9.1 %, had diagnoses which were made only on the basis of serological investigation (RSK); also, this figure varied in different hospitals from 4.5 to 15 %.

Thus, light or atypical and obliterated forms of the disease are common for contemporary typhus. Clinical forms of typhus infections of the same type are, undoubtedly, encountered even in foci of epidemic typhus, which follows from the considerable number of positive RSK in the older age groups with negative typhus anamnesis and visibly demonstrated in our material.

Evidently the correctness of the suggestions of K. N. Tokorevich (1958), stating that "typhus, in the sense of varied expression and typical development in

individual disease cases, does not represent an isolation from the class of other acute infections." Here occurs the similarity with bacterial and virus infections in the appraisal of retrospectively-revealed immunological indices in the province of typhus infection in older population groups with negative anamnesis for typhus, and having suffered unfortunate bouts with this infection.

As L. V. Gromashevskiy correctly indicates, if "obliterated" forms were present in typhus, this would predetermine, in the positive sense, the possibility of "asymptomatic" forms of typhus disease, corresponding to virus-carrying. Since, as is now determined, "obliterated" forms are, undoubtedly, met in typhus diseases, the question of its "asymptomatic" forms is predetermined a priori.

Parallel with this, the possibility has been proven out by foreign investigators of the presence, in all types of the most important rickettsial diseases, of long-lived latent forms of infection in persons who had been ill with the pathogenic agents in lymph nodes (Parker; Smedley and colleagues; and Price).

Our material does not yet include direct observations of "asymptomatic" forms of typhus disease if two cases are not counted which were discovered by the workers of the Gor'kivskiy Institute of Microbiology and Epidemiology (G. I. Grenauss, and others, 1958) in family foci of sporadic typhus; also, in one healthy subject a positive RSK at 1:20 titer was discovered and hemagglutination at 1:4000 titer, and in the other -- a RSK at 1:4 titer and hemagglutination at a 1:1000 titer. It should be noted that hemagglutination in the titers noted is obtained only in active infection or in the early stages of convalescence. The authors who discovered the cases cited are not certain that the given subjects did not have a light attack prior to their investigation to which they paid no attention. Thus, if the given case exhibited no "asymptomatic" infection proper, then, at least a very light form of typhus was present which was experienced, though the person was not bedridden. It is impossible, however, to omit the very thorough examinations carried out with positive results in recent years in Rumania under the leadership of the prominent specialist on typhus, Prof. D. Ciubesco.

The Rumanian authors, well-acquainted by past experience with epidemic typhus, in recent years also

encountered the epidemiological paradoxes of sporadic typhus. In connection with this, in analogy with us, they carried out mass serological investigations in the unfortunate rayonny with the aid of specific RSK, concentrating especially on the search for obliterated and asymptomatic forms in the foci of sporadic and epidemic typhus.

The results which are of interest to us, of the researches of Prof D. Combiesco who headed the large collective of research participants, are summarized in two theses, which we quote from the original: "The application of RSK allowed the recognition of numerous hidden cases of typhus which play an important role in epidemic typhus, since these forms of the disease are in large measure responsible for the appearance of sporadic forms of the disease"; "With the aid of RSK the hidden epidemiological foci, which had passed for sporadic examples, were exposed as obvious epidemiological foci by numerous cases."

We present an example of the discovery in foci of "asymptomatic" forms of typhus, which were found serologically.

1. In a ten-year old case the diagnosis of typhus was belatedly determined by a RSK at a 1:512 titer. Seven persons were found in contact with the diseased, ranging in age from 14-18 to 36-48 years of age. On studying the, a positive RSK was found in six at titers of 1:128, 1:64, 1:28, 1:32, 1:256, and 1:16. The focus was given special attention. Following the isolation of the ill person, among the remainder, regardless of the RSK presence at diagnostic titers (1:16 and higher), illnesses were not found.

The example given illustrates the actual focus of typhus infection, which in the absence of serological investigations of persons surrounding the diseased, would qualify only as a sporadic case of typhus.

We present another example where a case of clinically-manifest and asymptomatic typhus was subject to a repeated serological investigation.

2. The ill person had the RSK: 25/VI - 1:1024; 22/VII - 1:1024; 30/VII - 1:1024; 4/X - 1:125.

The RSK of the person who had been in contact with the diseased and who had not fallen ill were: 2/VIII - 1:512; 29/VII - 1:1024; 4/X - 1:100; in April of the following year -- 1:128 (Combiesco, 1957).

The examples cited pertain to foci of sporadic typhus. Below we present the discovery of "asymptomatic" forms in a focus of epidemic typhus.

3. The focus in Kucha-Voda. During January and February an epidemic outburst of typhus occurred with the appearance of 25 cases of the disease, of which half were found in schoolchildren. In the beginning of March a laboratory investigation was made of 168 persons of those who were in contact with the diseased cases, of that number were 33 pupils, ranging in age from 7 to 14 years of age and 135 were persons out of school, from 15 years of age and up.

The results of the serological investigations of the entire group by RSK are presented in Table 4.

Thus, among 33 pupils 14 were found to have positive RSK in titers of the range of 1:16 - 1:2048. In connection with this the authors note that all the cases with a positive serological reaction that were uncovered remained healthy (Combienco, 1958).

From Combienco collected data (1958) in 48 epidemic foci (in the presence of a considerable infestation here, judging by the illustrated presented), with the aid of RSK 2108 trials of serums of persons who had been in contact with typhus were investigated; here, in 947 cases (44 %) RSK was found to be positive with titers in the range of 1:4 to 1:2048, with an age distribution of 9 % positive in a group of children up to six years of age, and to 53 % positive within the ages of 51-60. Serological findings are detailed in Table 5.

TABLE FIVE

Results of the Serological RSK Investigation in the Focus of an Epidemic Outbreak of Typhus

Groups investigated	RSK Titers								
	0	4	8	16	32	64	128	256	2048
Pupils 7-14 yrs old	2	9	8	3	2	6	2	1	---
Persons not in school who are over 15 yrs and older	13	38	49	27	3	1	2	1	1

TABLE FIVE

RESULTS OF RSK SERIOLOGICAL INVESTIGATION IN FOCI OF EPIDEMIC  
OUTBREAKS OF TYPHUS (According to the data of D. Comblesco)

Number Studied	TSK Titers										
	0	4	8	16	32	64	128	256	512	1024	2048
Absolute Figure	1126	259	261	233	116	17	14	6	3	1	2
Percent	56	12	12	11	5.68	0.88	0.68	0.24	0.14	0.04	0.09

Thus, to sum up, titers in the range of 1:16 to 1:2048 were encountered in 392 persons, i.e., in 18.5% of those investigated.

As D. Comblesco points out, in the majority of the ill (dans la majorite des cas) with a showing of positive RSK, the infection was not discovered by clinical observation. However, in 43 out of 46 cases studied with the aid of RSK the immediate sources of infection were found.

With the above-indicated data of the Rumanian authors, concerning the contact with foci of epidemic typhus in the presence of a considerable infestation, it is interesting to compare the analogous data of M. A. Kulemina and K. P. Kuznetsova (1958) on the serological investigation of healthy cases with negative anamnesis for typhus in foci of sporadic typhus in the presence of only isolated cases of cephalic pediculosis. These data are presented in summary form in Table 6.

As is seen from the data presented, contacts in the foci of sporadic typhus with minimum infestation, evidently, were likewise accompanied by the appearance of immunological indicators of typhus infection, but in considerably lesser degree in comparison with the foci of epidemic typhus.

Thus, the question of the asymptomatic forms of typhus from the data of the Rumanian investigators is quite solidly based and, also, there is pertinent to this question sporadic foci as well as epidemic typhus. In any case these data do (indirectly, but sufficiently

logically in agreement with our material also, that is presented in this survey) deserve the most intense attention and influence the execution of corresponding investigations.

TABLE SIX

Results of RSK Serological Investigations in  
Foci of Sporadic Typhus (according to the data  
of M. A. Kulemina and A. P. Kuznetsova)

Number inves- tigated in the foci	Positive RSK cases of these		RSK Titers			
	Absolute figure	Percentage	1:10- 1:20	1:20- 1:40	1:40- 1:80	1:80 and up
851	85	10	29	27	29	0

we will turn now to the additional characterist-  
ics of sporadic typhus.

The great majority of foreign investigators in Europe and America identify sporadic forms of typhus with Brill's disease, considering them as relapses of previously experienced typhus. The relapse nature of sporadic typhus is defended here by K. N. Tokarevich and S. G. Nosing.

In Rumania, D. Combesco and his colleagues, admitting the relapse nature of repetitive typhus, parallel with this predicts the primary origin of sporadic forms, pointing out that with the aid of RSK the relations between sporadic diseases (by the presence in the foci of hidden forms of the infection) may be shown.

In our opinion it would be a mistake to register the observed cases of sporadic typhus under the generalized term "typhus." At the same time it would be incorrect to limit the category "sporadic typhus" only to cases of repetitive typhus from the data of anamnesis. Evidently, all the individual cases of typhus that do not tie in with a known source of infection, regardless of their nature, should be viewed and registered as "sporadic typhus" of the corresponding Brill's disease, in official nomenclature, VCZ. The basis for what has been stated is the fact that according to the data of our investigators there is no spec-

1]ial difference in the clinic of repetitive and period-  
ically sporadic typhus. Evidently they are similar by  
their relatively low infectiousness.

It may be supposed that the cause of contempor-  
ary typhus, retaining a visual resemblance to Provachek's  
rickettsia, is characterized by less effective patho-  
genic agents. Under our conditions of socio-economic  
progress, typhus more and more takes on the character-  
istics of a waning infection.

According to the joint announcement of all com-  
petent investigators -- native and foreign -- reliable  
diagnosis of contemporary forms of typhus demands the  
necessary application of serological diagnostics in  
the form of agglutination reactions or, better, reac-  
tions of complement fixation with Provachek's rickett-  
sia antigen. As has previously been noted, on the av-  
erage about 10 % of those ill with sporadic typhus,  
even in conditions of hospital observation, are diag-  
nosed only by RSK; also, in a number of cases typhus  
was retrospectively diagnosed, after they had already  
left the hospital where they had been with other diag-  
noses.

Also, from the most recent data of the Rumanian  
authors "epidemiological connections between separate  
sporadic attacks of typhus may be exposed and clarified  
only with the aid of serological investigations of cor-  
responding contingents" (D. Combesco).

Thus, on the basis of the material presented  
above, two needs of principal importance may be formu-  
lated: 1) to legislate the nomenclature of "sporadic  
typhus" in the above-mentioned sense; 2) to facilitate  
and inculcate into wide practice serological diagnostics  
of typhus by contemporary methods, having organized the  
mass production of moderately priced antigens of high  
quality.

In conclusion, there should be emphasized the  
pressing need for the most thorough possible study of  
"residual" forms of typhus from contemporary approach-  
es and with the use of contemporary methods of immuno-  
logical diagnosis of typhus infection. With this, in  
connection with the observations of the Rumanian authors,  
special attention should be paid to the detection of ob-  
literated and asymptomatic forms of the infection in  
foci of typhus, for the clarification of their condi-  
tions of origin and epidemiological significance. Use  
of the hemagglutination reaction (HAR) in parallel  
with RSK lends a special perspective. This reaction

provides the possibility of the differentiation of active forms of typhus infection and their most immediate convalescences in contrast to RSK, which at low titers does not differentiate retrospectively diagnosed forms from fresh ones (V. A. Yablonskaya, L. N. Astakhova, M. V. Matveyeva).

As the experiment of the application of HAR in typhus in comparison with its mass trial use among healthy contingents (over 4000 people investigated) showed, for the diagnostics of active forms of typhus infection HAR is conclusive at titers of 1:1000 and higher.

It is necessary to sever all ties with the widespread and completely unfounded conception of the imaginary mastery of the epidemiology of typhus.

It is necessary to make a few statements concerning "caution" heard in the school of conservative epidemiologists, as though the admission of obliterated and asymptomatic forms of typhus and the relapse theory of the origin of repetitive typhus is disorganizing and weakening the battle with typhus. To these warnings should be contrasted first of all, the fact that we are obligated in our conceptions to be subject to "Mr. Fact." On the other hand, the admission of the these indicated above are not contributing to the weakening but, on the contrary, to the strengthening of our means of combatting typhus. Sporadic and obliterated forms of typhus infection are potential sources of infection and in unfavorable conditions may produce flareups of epidemic typhus. Therefore, all the generally accepted methods for combatting typhus remain in force with the request that they be unconditionally implemented. Supplementing them, a request is presented for the wide application of contemporary immunological investigations, as in the clinic with the aims of diagnostics, thus to epidemiological investigations of foci of sporadic typhus. In other words, the system of methods to be taken, in the battle with the potential threat of typhus is not growing weaker but, on the contrary, is gaining in strength and is being armed with improved methods of work.

# BIBLIOGRAPHY

L. V. Gromashevskiy, in the book by L. V. Gromashevskiy and G. M. Vayndrakh, Chastnaya epidemiologiya (Local Epidemiology), Moscow, 1947, page 413.

P. F. Zdrovskiy and Ye. M. Golinevich, Ucheniye o rickettsiyakh i rickettsiozakh (Studies on Rickettsia and Rickettsial Diseases), Moscow, 1956.

K. N. Tokarevich, Rickettsiozy (Rickettsial Diseases), Leningrad, 1958.

D. Combiesco, I. Buzdugan, V. Binzar, et al, Arch. Roum. Path. Exer., 1957, 1958, Vol 26, page 185.

D. Combiesco, G. Zerneu, I. Buzdugan, et al, Arch. Inst. Pasteur Tunis, 1959, Vol 36, page 401.